

Distinguishing between enamel fluorosis and other enamel defects in permanent teeth of children

Aira Sabokseir, Ali Golkari, Aubrey Sheiham

Background: The inconsistent prevalence of fluorosis for a given level of fluoride in drinking water suggests developmental defects of enamel (DDEs) other than fluorosis were being misdiagnosed as fluorosis. The imprecise definition and subjective perception of fluorosis indices could result in misdiagnosis of dental fluorosis. This study was conducted to distinguish genuine fluorosis from fluorosis-resembling defects that could have adverse health-related events as a cause using Early Childhood Events Life-grid method (ECEL).

Methods: A study was conducted on 400 9-year-old children from areas with high, optimal and low levels of fluoride in the drinking water of Fars province, Iran. Fluorosis cases were diagnosed on the standardized one view photographs of the anterior teeth using Dean's and TF Indices by calibrated dentists. Agreements between examiners were tested. Early childhood health-related data collected retrospectively by ECEL method were matched with the position of enamel defects.

Results: Using both Dean and TF indices three out of four dentists diagnosed that 31.3% (115) children had fluorosis. After matching health-related events in the 115 (31.3%) of children diagnosed with fluorosis, 31 (8.4%) of children had fluorosis which could be matched with their adverse health-related events. That suggests that what was diagnosed as fluorosis were DDEs which may resemble fluorosis.

Discussion: The frequently used measures of fluorosis appear to overscore fluorosis. Use of ECEL method to consider health related events relevant to DDEs could help to differentiate between genuine fluorosis and fluorosis-resembling defects.

1 **Distinguishing between enamel fluorosis and other enamel defects in permanent**
2 **teeth of children**

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13 **Abstract**

14 **Background:** The inconsistent prevalence of fluorosis for a given level of fluoride in
15 drinking water suggests developmental defects of enamel (DDEs) other than fluorosis
16 were being misdiagnosed as fluorosis. The imprecise definition and subjective
17 perception of fluorosis indices could result in misdiagnosis of dental fluorosis. This study
18 was conducted to distinguish genuine fluorosis from fluorosis-resembling defects that
19 could have adverse health-related events as a cause using Early Childhood Events Life-
20 grid method (ECEL).

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23 cases were diagnosed on the standardized one view photographs of the anterior teeth
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25 were tested. Early childhood health-related data collected retrospectively by ECEL
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27 **Results:** Using both Dean and TF indices three out of four dentists diagnosed that
28 31.3% (115) children had fluorosis. After matching health-related events in the 115
29 (31.3%) of children diagnosed with fluorosis, 31 (8.4%) of children had fluorosis which
30 could be matched with their adverse health-related events. That suggests that what was
31 diagnosed as fluorosis were DDEs which may resemble fluorosis.

32 **Discussion:** The frequently used measures of fluorosis appear to overscore fluorosis.
33 Use of ECEL method to consider health related events relevant to DDEs could help to
34 differentiate between genuine fluorosis and fluorosis-resembling defects.

36 Introduction:

37 Despite the widescale use of well documented indices of dental fluorosis (Dean 1942;
38 Fejerskov 1988), when using them there is inconsistency in the prevalence rates of
39 fluorosis for a given level of fluoride in drinking water. The inconsistency in reporting
40 fluorosis occurs due to subjective perception of fluorosis by examiners. Therefore, there
41 is a strong possibility that other Developmental Defects of Enamel (DDE) rather than
42 excess intake of fluoride are being misdiagnosed as fluorosis (Atar & Körperich 2010).

43 The two widely used indices of dental fluorosis are Dean's Index (Dean 1934; Dean
44 1942) and the TF Index (TFI) (Thylstrup & Fejerskov 1978; Fejerskov 1988). None of
45 them clearly distinguish between defects caused by fluorosis and caused by other
46 factors. The differences between some of the diagnostic categories are uncertain,
47 vague, or insensitive. In Dean's Index, each individual is given one score, as a score for
48 the whole mouth, according to the two teeth most affected by fluorosis. Thylstrup and
49 Fejerskov reformulated Dean' Index as the TF Index (TFI). They used both clinical and
50 histological appearance of fluorosis and created a single coded index from 0 (normal) to
51 9 (Thylstrup & Fejerskov 1978). The TFI was modified and finalized in 1988 (Fejerskov
52 1988). Still the differentiation between some categories of TFI is not sufficiently precise.

53 As the main indices for diagnosing dental fluorosis include definitions that are imprecise,
54 it is not surprising that in some studies the reported prevalence of fluorosis was similar
55 despite the levels of fluoride in the drinking water being similar or different. For example,
56 a study in KwaNdebele (Africa) revealed that the prevalence of fluorosis was similar in
57 residents of areas with considerably different levels of fluoride in the water supply
58 (Lewis & Chikte 1995). A study in Andhra Pradesh (India) of four different areas with
59 different levels of fluoride in drinking water (< 0.7, 0.7-1.2, 1.3-4.0, >4 ppm) reported
60 100% dental fluorosis even in areas with optimum level of fluoride (Sudhir et al. 2009).
61 In Hong Kong, where the fluoride concentration of public water supplies was increased
62 from 0.5 ppm to 0.7 ppm, and then 1 ppm, the prevalence of DDE of maxillary incisors
63 decreased significantly with increases in the fluoride levels; from 92%, to 55%, and then
64 35% (Wong et al. 2006).

65 These inconsistent results suggest that DDEs other than fluorosis were being
66 misdiagnosed as fluorosis. Fluorosis and some DDEs are similar in appearance (Small
67 & Murray 1978; Pindborg 1982). Numerous local and systemic risk factors cause DDEs
68 (Small & Murray 1978; Pindborg 1982; Atar & Körperich 2010). Some DDEs may be of
69 genetic origin (Thesleff 2000; Atar & Körperich 2010) or caused by malnutrition or
70 diseases that occurred during early childhood (Pindborg 1982; Atar & Körperich 2010).
71 Using teeth as markers of significant health related events and considering these risk
72 factors could help to distinguish between genuine fluorosis and fluorosis resembling
73 DDEs.

74 Genuine fluorosis can best be distinguished from other DDEs by relating a DDE to
75 particular life events, such as a significant health related event. Such events can be
76 reasonably recorded using the life-grid method that helps people remember past events
77 more accurately (Blane 1996). The life-grid method has been widely used and been
78 very successful in obtaining the precise timing of past events in both qualitative and
79 quantitative studies (Berney & Blane 1997; Holland et al. 2000; Bell 2005). A specially
80 designed life-grid for early childhood containing developmental milestones and shorter
81 periods of time for the earlier years, is the Early Childhood Events Life-grid method
82 (ECEL) (Golkari 2009).

83 As there is a need for accurate data of dental fluorosis, the inconsistency of reports on
84 the prevalence of fluorosis suggests that more precise definitions and diagnostic
85 methods are needed for diagnosing dental fluorosis and distinguishing enamel fluorosis
86 from other DDEs. Therefore a study was planned with the objective of distinguishing
87 genuine fluorosis from fluorosis-resembling defects that could have adverse health-
88 related events as a cause, instead of just excess fluoride intake.

89

90 **Materials and methods:**

91 This cross-sectional study was performed on 400 9-year-old children of Fars province,
92 Iran. The children were randomly selected from areas of Iran with high, optimal and low
93 levels of fluoride in the drinking water. Fluorosis cases were diagnosed using Dean's
94 Index and TF Index (Dean 1942; Fejerskov 1988) by calibrated dentists. To differentiate
95 between genuine fluorosis and fluorosis resembling defects, early childhood data were
96 collected retrospectively by the ECEL method (Golkari 2009).

97 Ethical permission was obtained from the Ethical Committee of Shiraz University of
98 Medical Sciences (SUMS) and the Educational Head Office of the Fars Province. A
99 written consent form explaining the objectives and the stages of the study were sent to
100 the parents of the selected children. They were included only if the parents agreed to
101 take part.

102 The level of fluoride in water of each town of the province was obtained from their
103 primary health care trust. Towns were categorized into one of three fluoride categories;
104 high (>2.0 ppm), optimal (0.7- 1.0 ppm) and low fluoride (<0.4 ppm). One town was
105 selected randomly from each category. The selected towns were Gerash with high
106 fluoride (2.12- 2.85) in water, Sepidan with low fluoride (0.24- 0.29) in water, and Shiraz
107 including four zones was the city with an optimal range of fluoride (0.62- 1.22) in water .

108 Sample size was calculated based on the estimated prevalence of fluorosis in Iran
109 (61%), $d=6.1\%$ (10% expected prevalence), $\alpha=0.05$. As a result, a sample size of 246
110 was needed if a simple randomized sampling was used in all areas. The selection of
111 children in Gerash and Sepidan (with high and low fluoride levels) was done by simple
112 randomized sampling. However, in Shiraz (chosen as area with optimum fluoride level),
113 the simple randomization was not possible as it was a big city. Therefore, a stratified
114 randomization method had to be adopted. It was done using the four different
115 educational zones of the city. The number of required sample in this area was multiplied
116 by 2 ($k=2$) to increase the accuracy in the stratified sampling method that was used. The
117 response rate was suggested to be around 80%. Therefore, at the end, 100 9-year-old
118 schoolchildren of Gerash, 100 from Sepidan and 200 from Shiraz were selected from
119 lists of students obtained from Educational Head Office of each town.

120 Children aged from 9 years, 0 month to 9 years and 11 months who returned the signed
121 consent form were included in the study. Children were excluded if they had lived for
122 more than six months from birth to 5 years of age in other towns. Those who had less
123 than 7 permanent incisor teeth, those with orthodontic brackets, overlapping teeth, and
124 large restorations or severe extrinsic stains on their incisors were also excluded.

125 Intra-oral examinations were carried out to select children conforming to the study
126 criteria. The examinations were performed using a headlight, disposable mirrors and
127 tongue blades with children seated on a chair. Photographs of the dentition of selected
128 children were taken for the diagnosis of dental fluorosis. A one-view photograph was
129 taken of the anterior part of dentition using a digital camera (Nikon D7100, AF-S VR
130 Micro-Nikkor 105mm f/ 2.8 G IF-ED) based on methods described by Wong et al.
131 (2005). Children were asked to close their incisor teeth edge to edge. Cheeks and lips
132 were retracted so that all anterior teeth and some parts of upper and lower gums were
133 visible. The camera was adjusted to 15 degrees above the perpendicular to the central
134 incisors' plane to minimize specula reflection and burn outs (Ellwood et al. 1996).
135 Immediately after taking each photograph, it was assessed to confirm its quality, and
136 was repeated if necessary.

137 During the fluorosis assessment phase, first, the photographs were randomly ordered to
138 prevent bias induced by the assessors' foreknowledge of the fluoride in the area.
139 Photographs were then assessed by eight calibrated dentists who were blind to the
140 clinical condition and town of residence of the subjects. Four calibrated dentists
141 assessed the photographs based on Dean's Index, and another four used the TF Index
142 (Dean 1942; Fejerskov 1988). All dentists observed the photographs on one computer
143 with identical settings. The diagnosis of fluorosis was confirmed only if three out of four
144 dentists of each group agreed.

145 The objective of the next stage of study was to obtain data on early childhood adverse
146 events. The parents of all children participating in the study were invited to school for
147 interview. Name and date of birth of each child were double checked with their parents.
148 Parents were interviewed using the ECEL method (Golkari 2009). Information regarding

149 gestational age (preterm, term, delayed), birth weight, number of births, type of delivery
150 (natural, caesarean or facilitated delivery), trauma to baby during birth, and newborn
151 vitality score was obtained. Questions were asked relating to personal life line,
152 residential status, occupation of parents/guardians, and child activity line which could
153 help parents to remember adverse health-related events. Afterwards, any illnesses the
154 child had suffered were recorded. For each illness the parent was asked about the
155 name or description of illness, age at which the illness started, duration, perception of
156 severity (mild, moderate, or severe), if went to doctor, medication if used, and
157 hospitalization. Information regarding hospitalizations (age, duration, reason, type of
158 anesthesia if used, and name of hospital), and falls and accidents (age, cause, trauma
159 to face or teeth, hospitalization, and breathing status right after accident as an indicator
160 of severity well known by parents) were also obtained.

161 The timing of childhood adverse health-related conditions was matched with the timing
162 of formation and calcification of each part of permanent incisors (Golkari 2009). If no
163 adverse life condition could be matched to the position of a defect diagnosed as
164 fluorosis, the case was considered as genuine fluorosis. However, if a health-related
165 adverse condition could be matched to an enamel defect diagnosed as fluorosis, the
166 case was considered as fluorosis-resembling defect. Using this method, diagnosed
167 fluorosis defects were divided into genuine fluorosis and fluorosis resembling defects.

168 SPSS software (version 22) was used for data analysis. Agreements between
169 examiners who assessed fluorosis using Dean's Index and TF Index were tested using
170 Kappa-coefficient. McNemar, and Pearson correlation tests were also used to compare
171 the results reported by each pair of dentists. The prevalence of fluorosis in the three
172 selected areas was compared by one-way ANOVA. The relationship of fluorosis with the
173 level of fluoride in water, and sex was tested using Logistic regression. Childhood
174 adverse health-related conditions were matched to enamel defects diagnosed as
175 fluorosis one by one.

176

177 **Results:**

178 The number of 9-year-old children included in the study was 376; 171 (46%) girls and
179 196 (53 %) boys. The number of included children from Gerash (high F), Shiraz (optimal
180 F) and Sepidan (low F) were 88, 189, and 90 respectively .

181 Using both Dean and TF indices, three out of four dentists diagnosed that 31.3% (115)
182 of children had fluorosis. The percentage of fluorosis cases in areas with high, optimal,
183 and low range of fluoride in water was 58.0%, 29.1%, and 10.0% respectively
184 ($p < 0.001$). Logistic regression showed that there was a positive relationship between
185 fluorosis and fluoride in the drinking water ($p < 0.001$). There was no relationship
186 between fluorosis and children's sex ($p = 0.228$).

187 There were significant differences among dentists who scored photographs using
188 Dean's and TF indices. Among the four dentists who assessed photographs according
189 to Dean's Index, the difference in the number of cases diagnosed as fluorosis was
190 statistically different between each two dentists ($p < 0.001$). There was only a slight
191 (kappa was between 0 and 0.2) or fair (kappa was between 0.2 and 0.4) agreement
192 between them (Table 1). Similar results of agreement were observed among the four
193 dentists who scored children using the TF Index. Although there was not a high
194 agreement among dentists, a positive correlation was observed ($p < 0.001$) (Table 2).

195 Adverse health-related events were found in 84.7% (311) of children of whom 8.4% of
196 children had fluorosis-resembling defects. After matching health-related events in the
197 115 (31.3%) of children diagnosed with fluorosis, 31 (8.4%) of children had fluorosis
198 which could be matched with their adverse health-related events. That suggests that
199 what was diagnosed as fluorosis were DDEs which may resemble fluorosis. Therefore,
200 it was concluded that the percentage with genuine fluorosis was 22.9% and not 31.3%.
201 The fluorosis resembling defects (8.4% of all children) included 26.9% of what was
202 diagnosed as fluorosis by Dean's and TF Indices. The percentage of genuine fluorosis
203 in areas with high, optimal and low levels of fluoride in water was 47.7%, 20.6%, and

204 3.3% respectively. The percentage of fluorosis-resembling defects in areas with high,
205 optimal, and low range of fluoride in water was 10.2%, 8.5%, 6.7% respectively.

206

207 **Discussion:**

208 Fluoride is one of the most successful measures for prevention of dental caries in public
209 health (Petersen & Lennon 2004). However, there has always been controversy about
210 using fluoride because of fluorosis (Sapolsky 1968; Null & Feldman 2003; Ananian et al.
211 2005). Reports of a high prevalence of fluorosis in communities have led to objections
212 to fluoride. Therefore, there is a need for a precise way to diagnose dental fluorosis.
213 Many local and systemic risk factors cause DDEs. Some DDEs are similar to enamel
214 fluorosis and should be differentiated from genuine fluorosis. The difference has not yet
215 been evaluated. The main objective of this study was to distinguish fluorosis that
216 resembled DDEs caused by adverse health-related events, and not caused by excess
217 fluoride intake.

218 A systematic review reported that the prevalence of fluorosis in Iran was 61%.
219 Considering the average concentration of fluoride in water in Iran was 0.43 ± 0.17
220 (Azami-Aghdash et al. 2013), the reported prevalence of fluorosis was high and
221 questionable. The inconsistency of the prevalence of fluorosis with the level of fluoride
222 in water reported in that study and in many other different parts of the world (Lo &
223 Bagramian 1996; Sudhir et al. 2009; Arif et al. 2013) suggests that dental fluorosis was
224 misdiagnosed.

225 The overall prevalence of diagnosed dental fluorosis in the current study was 31.3%.
226 However, by using the ECEL method and considering health-related events, the
227 prevalence of genuine dental fluorosis was 22.6% as there were 8.4% with fluorosis
228 resembling defects. That illustrates that fluorosis could be ruled out as the main cause
229 of about 27% of what was incorrectly diagnosed as fluorosis.

230 The current study also showed significant differences among the dentists who scored
231 photographs to diagnose fluorosis according to Dean's and TF Indices ($p < 0.001$). This
232 finding indicates that both Dean and TF Indices are too subjective and therefore not
233 precise ways to diagnose dental fluorosis.

234 These two Indices could lead to misdiagnosis of fluorosis resembling defects as
235 genuine fluorosis. Tavener et al also concluded that interpretation of criteria could be
236 different among examiners and stressed the necessity for standard methods to measure
237 dental fluorosis (Tavener et al. 2007). Some studies have shown good to excellent
238 agreement among examiners (Kumar et al. 2000). However, comparing the methods,
239 the lack of bias was an advantage of the current study, as photographs were assessed
240 instead of clinical examinations.

241 This study indicates that by considering adverse health-related events, it is possible to
242 distinguish genuine fluorosis from fluorosis-resembling defects. When the timing of an
243 adverse condition matches the timing of development of the part of enamel defect, the
244 adverse event could be the cause of the defect, not fluoride. If no adverse condition
245 could be matched to a defect, excess fluoride could, with caution, be considered as the
246 cause. On the other hand, if fluoride is considered the cause of defects, such defects
247 should not be seen on the teeth of children who experienced the same adverse health-
248 related conditions and were living in areas with low levels of fluoride in water. In case of
249 generalized defects, fluoride, genetic, or severe underlying systematic disease, either
250 individually or as a combination could be the cause.

251 One limitation of this study is that even if an adverse health-related event could be
252 exactly matched to a fluoride-resembling defect in terms of time and place, it could not
253 be definitely considered that the adverse event was the cause of the defect and fluoride
254 was not the cause. On the other hand, if an adverse condition could not be matched to
255 a supposedly fluorosis defect, one could not be sure that fluoride was the cause. In fact,
256 the definite diagnosis of fluorosis is not possible by this method and the only exact way
257 to diagnose dental fluorosis would be by microscopic or chemical analysis. However,

258 although the ECEL method cannot prove that fluoride is or is not the cause of defect, it
259 can help in differentiate between genuine fluorosis and fluorosis-resembling defects.

260 The diagnosis of fluorosis is more complicated than acknowledged. Existing fluorosis
261 indices could lead to misjudgment about using fluoride. However, the ECEL method to
262 record health-related life events is a promising method to help differentiate between
263 genuine fluorosis and fluorosis-resembling defects.

264 **Conclusion:**

265 Fluorosis indices, if used alone, could result in misdiagnosis of dental fluorosis.
266 Information about adverse health-related conditions linked to DDEs at specific positions
267 on teeth could help to differentiate between genuine fluorosis and fluorosis-resembling
268 defects.

269

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276

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333

335 Table1. The comparison of fluorosis scores according to Dean's Index by four calibrated
 336 dentists

	Test	Examiner 2	Examiner 3	Examiner 4
Examiner 1	Kappa value	0.16	0.16	0.07
	McNemar p-value	<0.001	<0.001	<0.001
	Correlation coefficient	0.528	0.454	0.458
Examiner 2	Kappa value		0.34	0.37
	McNemar p-value		0.125	<0.001
	Correlation coefficient		0.584	0.575
Examiner 3	Kappa value			0.29
	McNemar p-value			0.003
	Correlation coefficient			0.614

337

339 Table 2.The comparison of fluorosis scores according to TF Index by four calibrated
 340 dentists

	Test	Examiner 2	Examiner 3	Examiner 4
Examiner 1	Kappa value	0.06	0.21	0.34
	McNemar p-value	<0.001	<0.001	<0.001
	Correlation coefficient	0.381	0.465	0.498
Examiner 2	Kappa value		0.27	0.18
	McNemar p-value		<0.001	<0.001
	Correlation coefficient		0.472	0.503
Examiner 3	Kappa value			0.36
	McNemar p-value			<0.001
	Correlation coefficient			0.539

341